(FILE 'HOME' ENTERED AT 16:43:26 ON 02 JAN 2003)

FILE 'MEDLINE, EMBASE, CANCERLIT, BIOTECHDS, CAPLUS' ENTERED AT 16:44:05 ON 02 JAN 2003 316995 S ACRYLIC OR METHACRYLIC ACID OR MALEIC ANHYDRIDE OR EMA L1255 S EQUINE INFLUENZA VIRUS L2 L3 2 S L1 AND L2 2 DUP REM L3 (0 DUPLICATES REMOVED) L4L5361873 S VACCINE OR ADJUVANT L6 473 S L5 AND L1 L72184896 S PLASMID OR DNA rs17 S L6 AND L7 15 DUP REM L8 (2 DUPLICATES REMOVED) L9 2 S L1 AND DNA VACCINE L10 L11 5163 S DNA VACCINE 5 S L11 AND (POLYMER AND COPOLYMER) L12 4 DUP REM L12 (1 DUPLICATE REMOVED) L13 158945 S ADJUVANT L14376 S L14 AND L1 L15 L16 12 S L15 AND L7 L17 11 DUP REM L16 (1 DUPLICATE REMOVED) L18 648 S L11 AND L14 1278238 S POLYMER OR MICROPARTICLE OR COPOLYMER OR POLYMERIC L19 28 S L19 AND L18 L20 L21 20 DUP REM L20 (8 DUPLICATES REMOVED) 172 S L1 AND VECTOR L22 L23 54 S L22 AND (DNA OR PLASMID)

42 DUP REM L23 (12 DUPLICATES REMOVED)

L24

ANSWER 6 OF 11 BIOTECHDS COPYRIGHT 2003 THOMSON DERWENT AND ISI L17 ΑN 2000-02300 BIOTECHDS Live recombinant vaccine comprising virus vector and polymeric ΤI adjuvant, particularly directed against animal herpes and influenza viruses; recombinant vaccine production with a virus vector encoding a pathogen gene and an adjuvant for virus infection vaccination ΑU Audonnet J C; Minke J M PA Merial LO Lyons, France. PΙ WO 9944633 10 Sep 1999 ΑI WO 1999-FR453 1 Mar 1999 FR 1998-2800 3 Mar 1998 PRAI DТ Patent French LΑ WPI: 2000-022918 [02] OS A live recombinant vaccine which consists of a virus vector (A) AΒ containing a heterologous DNA sequence (I) (particularly encoding a gene from a pathogen) and at least on adjuvant (II), i.e. a methylacrylic acid polymer or a copolymer of maleic anhydride and alkenyl derivatives, is new. Also claimed is a vaccination kit which consists of lyophilized (A) and a solution of (II), in separate containers. These new vaccines may be particularly useful for protecting against animal herpes and influenza viruses, but they may also be useful for protecting against cat leukemia, tetanus toxin and dog distemper. The vaccines may be administered either parentally via a s.b., i.m. or i.d. injection, or mucosally. In an example, virus RNA from horse influenza virus strain Prague 56 was cloned into a plasmid to form vector plasmid pJT008. pJT008 was then linearized and used for in vitro recombination with a commercial

canary-pox virus to form vCP1502 recombinant virus. (40pp)

ANSWER 6 OF 11 BIOTECHDS COPYRIGHT 2003 THOMSON DERWENT AND ISI L17 AN 2000-02300 BIOTECHDS Live recombinant vaccine comprising virus vector and polymeric ΤI adjuvant, particularly directed against animal herpes and influenza viruses; recombinant vaccine production with a virus vector encoding a pathogen gene and an adjuvant for virus infection vaccination Audonnet J C; Minke J M ΑU PA Merial LO Lyons, France. PΙ WO 9944633 10 Sep 1999 ΑI WO 1999-FR453 1 Mar 1999 FR 1998-2800 3 Mar 1998 PRAI DΤ Patent French T.A WPI: 2000-022918 [02] OS A live recombinant vaccine which consists of a virus vector (A) AB containing a heterologous DNA sequence (I) (particularly encoding a gene from a pathogen) and at least on adjuvant (II), i.e. a methylacrylic acid polymer or a copolymer of maleic anhydride and alkenyl derivatives, is new. Also claimed is a vaccination kit which consists of lyophilized (A) and a solution of (II), in separate containers. These new vaccines may be particularly useful for protecting against animal herpes and influenza viruses, but they may also be useful for protecting against cat leukemia, tetanus toxin and dog distemper. The vaccines may be administered either parentally via a s.b., i.m. or i.d. injection, or mucosally. In an example, virus RNA from horse influenza virus strain Prague 56 was cloned into a plasmid to form vector plasmid pJT008. pJT008 was then linearized and used for in vitro recombination with a commercial

canary-pox virus to form vCP1502 recombinant virus. (40pp)

ANSWER 5 OF 11 BIOTECHDS COPYRIGHT 2003 THOMSON DERWENT AND ISI L17 AN 2000-00707 BIOTECHDS Vaccine containing naked DNA and acrylic acid polymer TI or maleric anhydride copolymer, for protection against virus or bacterial diseases in animals; vector plasmid-mediated gene transfer and expression in horse, pig, cattle, bird, dog or cat as a nucleic acid vaccine for bacterium and virus infection therapy Audonnet J C F; Minke J M ΑU PA Merial LO France. PΙ FR 2776928 8 Oct 1999 ΑI FR 1998-4409 3 Apr 1998 FR 1998-4409 3 Apr 1998 PRAI DTPatent French LΑ WPI: 1999-593389 [51] OS A nucleic acid vaccine which consists of naked DNA that AΒ includes and expresses in vivo, a sequence (I) which encodes an antigenic protein and at least one adjuvant (III) that is an acrylic or methacrylic acid polymer or a copolymer of maleric anhydride with an alkenyl derivatives, is new. claimed is a method for using (III) as an adjuvant in a nucleic acid vaccine containing, and expressing in vivo a heterologous sequence. These new nucleic acid vaccines may be useful for protecting animals (pigs, horses, dogs, cattle, cats and birds) against a wide variety of virus and bacterial infections. The vaccines have the advantage of being simple and easy to prepare (simply by mixing components) and they do not involve any strong interactions between DNA and other components that are likely to cause complex formation. In an example, reverse-transcription polymerase chain reaction was used to construct 3

plasmid vectors which were used as the nucleic acid vaccines.

(33pp)

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ANSWER 7 OF 11 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1999:659270 CAPLUS
DN
     131:298650
ΤI
     Polymer adjuvants for use with vector vaccines
     Audonnet, Jean-christophe Francis; Minke, Jules Maarten
IN
PA
     Merial, Fr.
     PCT Int. Appl., 37 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LΑ
     French
FAN.CNT 1
     PATENT NO.
                     KIND
                           DATE
                                           APPLICATION NO.
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                                                         19990322
PΙ
     WO 9951269
                     A1
                           19991014
                                          WO 1999-FR666
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             DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
             JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
             TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     FR 2776928
                      Α1
                            19991008
                                           FR 1998-4409
                                                            19980403
     FR 2776928
                            20000623
                       В1
     CA 2327389
                                           CA 1999-2327389
                            19991014
                                                            19990322
                      AΑ
    AU 9928448
                            19991025
                                           AU 1999-28448
                      Α1
                                                            19990322
    AU 744964
                      В2
                            20020307
     BR 9909342
                      Α
                            20001212
                                           BR 1999-9342
                                                            19990322
     EP 1066055
                                           EP 1999-909069
                      Α1
                            20010110
                                                            19990322
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 2002510651
                       T2
                            20020409
                                           JP 2000-542039
                                                            19990322
PRAI FR 1998-4409
                       Α
                            19980403
     WO 1999-FR666
                      W
                            19990322
     Polymer adjuvants that increase the efficacy of vector vaccines carrying
AB
     an expression cassette for an antigen gene of a pathogen are described.
     The polymers are acrylic or methacrylic polymers and the
    maleic anhydride copolymers and alkenyl deriv. The
     adjuvant compd. is preferably a carbomer or an EMA.RTM..
     Construction of expression vectors for a no. viral antigen genes were
     constructed using the com. expression vector pVR1012 is described.
     Inoculation of horses, swine, cattle, and dogs with these vectors with
     Carbopol 974P as an adjuvant is demonstrated. Use of the
     adjuvant led to the appearance of antibody to the antigens.
RE.CNT 3
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
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ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 18 OF 20 MEDLINE DUPLICATE 3

AN 1998382497 MEDLINE

DN 98382497 PubMed ID: 9714699

- TI Intranasal administration of HIV-DNA vaccine formulated with a polymer, carboxymethylcellulose, augments mucosal antibody production and cell-mediated immune response.
- AU Hamajima K; Sasaki S; Fukushima J; Kaneko T; Xin K Q; Kudoh I; Okuda K
- CS Department of Bacteriology, Yokohama City University School of Medicine, Yokohama, 236, Japan.
- SO CLINICAL IMMUNOLOGY AND IMMUNOPATHOLOGY, (1998 Aug) 88 (2) 205-10. Journal code: 0356637. ISSN: 0090-1229.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals; AIDS
- EM 199809
- ED Entered STN: 19981008 Last Updated on STN: 19981008 Entered Medline: 19980929
- AΒ We previously reported that intramuscular (i.m.) immunization of DNA vaccine encoding human immunodeficiency virus type 1 (HIV-1)IIIB env and rev genes alone or in combination with appropriate adjuvant induces substantial and enhanced immune response against HIV-1. In the present study, we examined whether a polymer, low-viscosity carboxymethylcellulose sodium salt (CMCS-L), has an adjuvant effect on immune response induced by DNA vaccination. BALB/c mice were immunized with HIV-DNA vaccine formulated with CMCS-L via the intranasal (i.n.) and i.m. routes. The combination with the polymer elicited higher levels of antigen-specific serum IgG and fecal IgA antibodies than DNA vaccine alone. For cell-mediated immunity, HIV-specific delayed-type hypersensitivity response and cytotoxic T lymphocyte activity were measured by the footpad-swelling test and the 51Cr-release assay, respectively. Both were enhanced by the combination with CMCS-L via i.n. and i.m. inoculation. Cytokine analysis in culture media of bulk splenocytes harvested from immunized animals showed higher levels of IL-4 production in i.n. -immunized mice compared with i.m.-immunized mice. Nevertheless, the increased IFN-gamma production resulting from the combination with CMCS-L was observed only in i.n.-immunized mice. These data indicate that i.n. immunization of HIV-DNA vaccine formulated with CMCS-L enhances HIV-specific mucosal antibody (Ab) and systemic Ab and cell-mediated immune response. Copyright 1998 Academic Press.

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ANSWER 2 OF 20 CAPLUS COPYRIGHT 2003 ACS
L21
     2002:51240 CAPLUS
AN
     136:107525
DN
    Microspheres and adjuvants for DNA vaccine delivery
ΤI
     Johnson, Mark E.; Mossman, Sally; Cecil, Tricia; Evans, Lawrence
IN
PA
     Corixa Corporation, USA
so
     PCT Int. Appl., 60 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LΑ
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                            _____
                                          WO 2001-US21780 20010709
PI
     WO 2002003961
                      A1
                            20020117
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             FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG,
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             MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
             TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD,
             RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           AU 2001-71976
                                                            20010709
                      Α5
                            20020121
     AU 2001071976
                                           US 2001-901829
                                                            20010709
     US 2002032165
                       Α1
                            20020314
PRAI US 2000-216604P
                       P
                            20000707
     WO 2001-US21780
                       W
                            20010709
     A nucleic acid delivery system that offers, in one system, a combination
AΒ
     of high encapsulation efficiency, rapid release kinetics and preservation
     of DNA in a supercoiled form is provided. The nucleic delivery system
     comprises nucleic acid mols., such as a DNA, encapsulated in biodegradable
     microspheres, and is particularly suited for delivery of DNA vaccines.
     The invention further provides an adjuvant for modulating the
     immunostimulatory efficacy of microspheres encapsulating nucleic acid
     mols. comprising an aminoalkyl glucosaminide 4-phosphate (AGP). Thus, a
     quick release, high efficiency, porous, 1-10 .mu.m DNA microsphere
     formulation was developed by using PLG copolymer and tested.
     Cytotoxic T-lymphocyte (CTL) responses to 2 antigens, Her-2/neu and TbH9,
     were generated using these DNA microspheres. I.m. and i.p. routes were
     the best for CTL elicitation. Several AGPs provided substantial CTL
     adjuvant activity to the DNA microspheres. Sodium.
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(16 ref) fluoride.

L24 ANSWER 22 OF 42 BIOTECHDS COPYRIGHT 2003 THOMSON DERWENT AND ISI ΑN 1998-07238 BIOTECHDS ΤI Production of nucleic acid conjugates; plasmid DNA and RNA conjugate preparation for use in gene transfer and sense, antisense gene expression control

ΑU Bayer E; Fritz H; Maier M SKW-Trostberg PA

LO

Trostberg, Germany.

PΙ DE 19746362 30 Apr 1998

DE 1997-1046362 21 Oct 1997 ΑI

PRAI DE 1997-1046362 21 Oct 1997

DT Patent

LΑ German

WPI: 1998-252414 [23] os

A new process for the production of conjugates of nucleic acids with AΒ polymer nanoparticles involves subjecting sparingly water-soluble vinylic monomers to emulsion polymerization in an ag. medium in the presence of a cationic radical initiator and in the absence of an emulsifier, preferably purifying the suspension by diafiltration or centrifugation, and reacting the resulting polymer suspension with a nucleic acid at 10-30 deg and pH less than 11. The conjugates are useful for gene transfer or for sense or antisense control of gene expression. Conjugates with high nucleic acid loadings and adequate resistance to enzyme degradation can be produced. The monomers preferably have a water solubility below 20 g/l and are selected from styrene, acrylic acid derivatives and methacrylic acid derivatives. The polymer suspension has a particle size of 10-1,000 nm. The nucleic acid is optionally chemically modified DNA or RNA with a length of 7-40 nucleotides, and is preferably a plasmid. (5pp)

- L24 ANSWER 11 OF 42 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.
- AN 2001213999 EMBASE
- TI Copolymers of amine methacrylate with poly(ethylene glycol) as vectors for gene therapy.
- AU Rungsardthong U.; Deshpande M.; Bailey L.; Vamvakaki M.; Armes S.P.; Garnett M.C.; Stolnik S.
- CS S. Stolnik, School of Pharmaceutical Sciences, University of Nottingham, University Park, Nottingham NG7 2RD, United Kingdom. snjezana.stolnik@nottingham.ac.uk
- SO Journal of Controlled Release, (12 Jul 2001) 73/2-3 (359-380). Refs: 44
  - ISSN: 0168-3659 CODEN: JCREEC
- PUI S 0168-3659(01)00295-4
- CY Netherlands
- DT Journal; Article
- FS 022 Human Genetics 037 Drug Literature Index
- LA English
- SL English
- A series of structurally related copolymers of tertiary amine methacrylate AB with poly(ethylene glycol) (PEG) were investigated for their potential to serve as vectors for gene therapy. The effects of copolymer structure on the complexation and transfection ability were assessed. The ability of the PEG-based copolymers and DMAEMA homopolymer to bind and condense DNA was confirmed by gel electrophoresis, ethidium bromide displacement and transmission electron microscopy. The presence of PEG in the copolymers had a beneficial effect on their ability to bind to DNA. Colloidally stable complexes were obtained for all the PEG-copolymer systems as shown by uniformly discrete spherical images from transmission electron microscopy and approximate diameters of 80-100 nm by dynamic light scattering studies. DMAEMA homopolymer, however, produced agglomerated particles, confirming the important role played by the PEG chains in producing compact stable DNA complexes. Assessment of the effect of ionic strength of the buffer on the complexation and dissociation of the complexes indicated the importance of both electrostatic and non-electrostatic interactions in the polymer-DNA complexation. In vitro transfection experiments showed that DMAEMA homopolymer gave the highest level of transfection comparable to a control poly-L-lysine (PLL) system. The PEG-based copolymers gave reduced levels of transfection, most likely due to the steric stabilization effect of a PEG corona. .COPYRGT. 2001 Elsevier Science B.V.